

Fading funds: US financial backing for high-energy physics has slowed down significantly.

fusion research programme in the United States lost one-third of its budget in 1995 after Congress decided that US involvement in the International Thermonuclear Experimental Reactor was too expensive to pursue.

Since the 1998 study, physicists at Fermilab and elsewhere have been developing the

case for a 'neutrino factory', which would generate an intense beam of neutrinos from the decay of muons circulating in a race-track-shaped ring.

Such an experiment, which would cost about a billion dollars, could build on current efforts in Japan, the United States and Europe to detect neutrino oscillation, and therefore measure the mass of the neutrino. Some advocates of the neutrino factory see it as a realistically priced, intermediate step towards a muon collider that could eventually be built at Fermilab.

"There are a lot of people worrying about their own survival, and they are not prepared to take the NLC on as a major thrust," says Lubell. One laboratory official goes further, saying that the current budget crunch "has created a crisis mode of thinking, close to a feeling of panic" in the field which, the official adds, is not conducive to reaching a consensus on future projects.

"It's a very difficult situation," says Nick Samios, former director of the Brookhaven National Laboratory. "Our credibility is not

high, because of the Superconducting Super Collider debacle." Samios supports the NLC construction, but notes that current plans for a collider with an initial energy of 500 GeV, upgradable to 1 TeV, falls well short of a previous assumption by US physicists that such a machine should aim for an energy of 1.5 TeV.

Whatever its eventual specification and site, US physicists are in agreement that the NLC will have to be a truly international project. The \$20 million annual research programme into the NLC, which is centred at SLAC, has been closely coordinated with a similar programme at the High Energy Accelerator Research Organization (KEK) in Japan.

At a meeting in London earlier this month, the global science forum of the Paris-based Organization for Economic Cooperation and Development discussed the formation of a high-energy physics workshop that could provide the necessary framework for the international construction of machines such as the NLC.

Colin Macilwain

'Saturation screen' lets zebrafish show their stripes

London

Over the next year, a mammoth project involving some 17 million tropical fish is expected to yield a treasure trove of data for researchers working on the genetics of vertebrate development.

Artemis Pharmaceuticals, a company based in Cologne, announced last week that it has joined forces with Christiane Nüsslein-Volhard of the Max Planck Institute for Developmental Biology in Tübingen to conduct a zebrafish 'saturation mutagenesis screen'.

The project, dubbed the Tübingen 2000 Screen, is already under way, and will cost around US\$10 million. Its goal is to apply chemical mutagens to zebrafish sperm to introduce point mutations randomly throughout the genome.

Researchers will breed fish using the treated sperm, and screen for those that show developmental abnormalities. From these animals, geneticists can then try to clone the mutated gene. Applying this technique on a sufficiently large scale should make it possible to mutate every important developmental gene — hence the term 'saturation' mutagenesis.

Nüsslein-Volhard shared a Nobel prize in 1995 for her work in developing saturation mutagenesis in the fruitfly *Drosophila*. By this time, she had also begun to apply the technique to the zebrafish (*Danio rerio*) to investigate the function of genes that control the development of vertebrates. This popular aquarium fish is ideal for

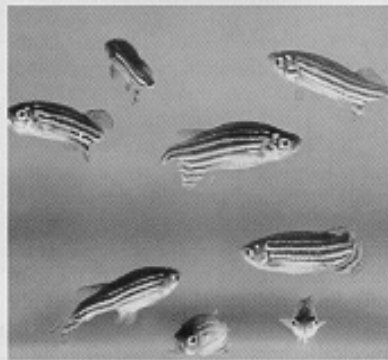
developmental studies, as its embryos are transparent, making it easy to spot abnormalities in internal structures.

Nüsslein-Volhard's previous zebrafish screen produced more than 1,000 mutants. A parallel screen — conducted by a team led by Wolfgang Driever, now at the University of Freiburg — produced half as many mutants again. Mutants from both screens were described in a special issue of the journal *Development* (see *Nature* 384, 514; 1996). "A lot of interesting mutants were found, and it has been possible to start to characterize some of those molecularly," says Phil Ingham, a developmental biologist at the University of Sheffield.

But the original screens failed to mutate every important zebrafish developmental gene. "We probably didn't get to saturation," says Nüsslein-Volhard, who is a co-founder of Artemis. In addition, some interesting mutants may have been overlooked because researchers didn't spot subtle developmental abnormalities in every single organ and physiological system.

This time, researchers with different expertise will examine the same fish to ensure that fewer mutants get overlooked. The screen will be twice the scale of Nüsslein-Volhard's previous project. Peter Stadler, chief executive officer of Artemis, says the project will employ about 50 people.

Scientists at Artemis will mainly screen for abnormalities giving information on gene function that could provide leads for drug development. These include



Variety shoal: zebrafish with the *mini fin* gene, one of a host of mutants known in the species.

abnormalities in the production of cartilage and bone, and in the cardiovascular system.

Nüsslein-Volhard's team, together with researchers from elsewhere in Germany, Britain and the United States, will look for mutants yielding fundamental insights into the processes of development. These will include fish with abnormalities in the formation of the eyes and the neural crest — paired strips of cells that form early in development, which give rise to a variety of tissues including the central nervous system.

Artemis will have six weeks to patent any discoveries. After this period, results will be made available to other scientists in the normal way, says Stadler.

Peter Aldhous

<http://www.artemis-pharmaceuticals.de/html/topic2000.html>

http://www.sb.tuebingen.mpg.de/abt3/home_text.html